WO 2004/043406

1/25

B

ATM

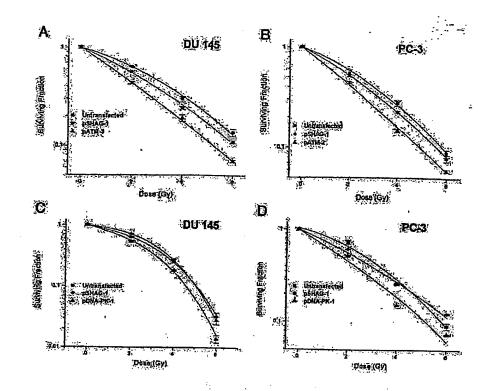
B Saktin

DNA-PK

G-actin

Translectani.

4 158 b 25 p 10 10 / 534010



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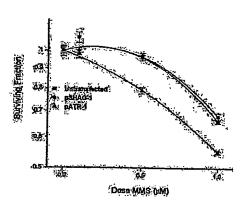
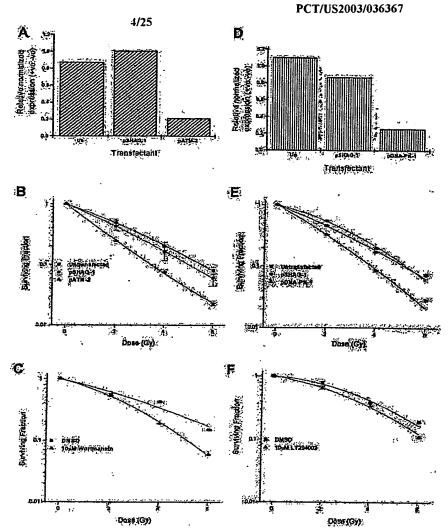
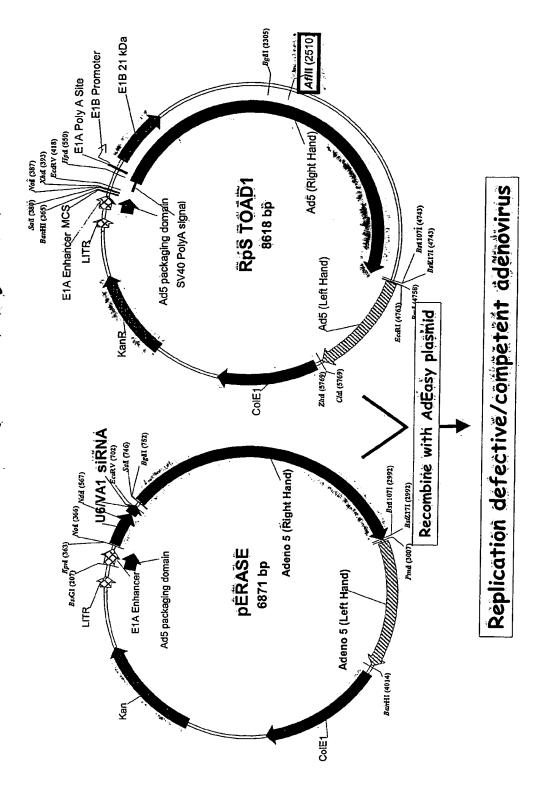


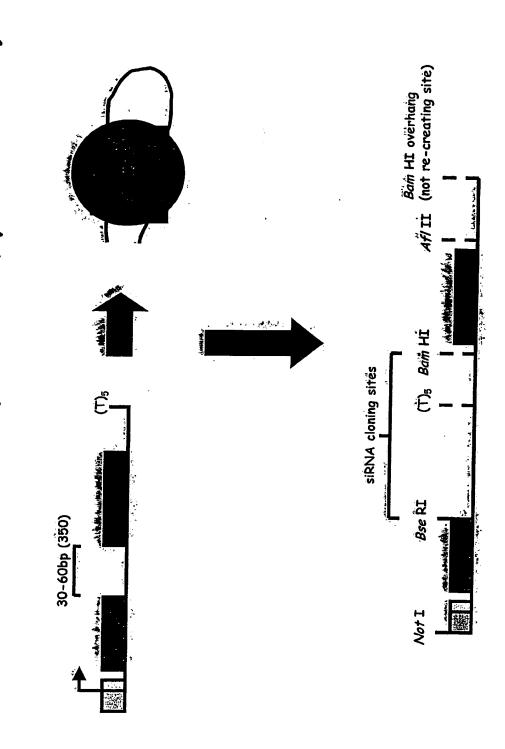
Fig. 4



# Ad5 siRNA (ERASE) vectors



An alternative pol III promoter (synthetic VA1)



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Effectiveness of VA1 promoter in PC-3 Luc cells

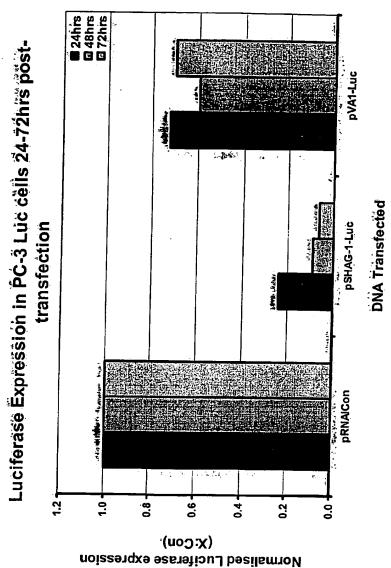
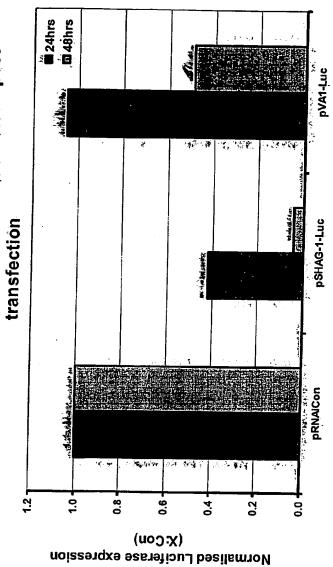


Fig. 7

**DNA Transfected** 

Effectiveness of VA1 promoter in 293 Luc cells

Luciferase Expression in 293 Luc cells 24-48hrs post-

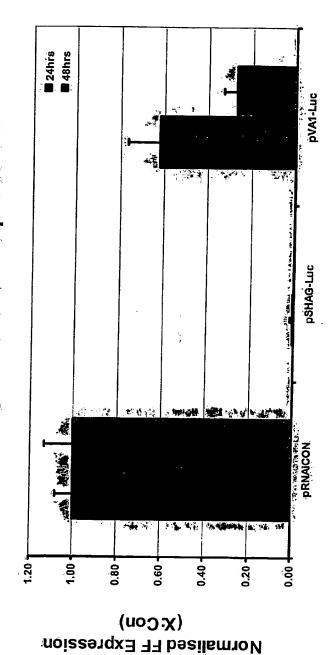


**DNA Transfected** 

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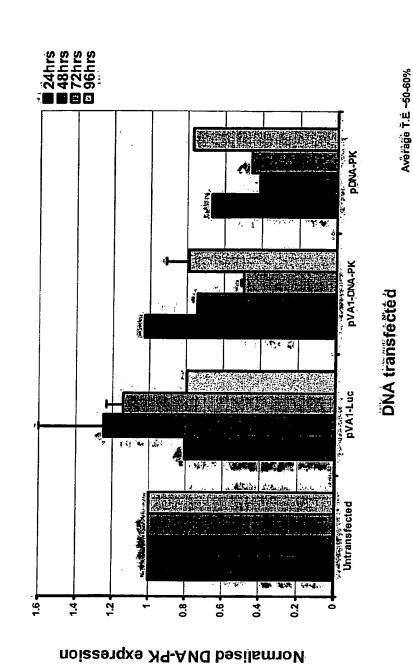
Effectiveness of VA1 promoter in DU 145 cells: co-transfection experiments

Fire Fly Luciferase Expression



VA1 versus U6-mediated down-regulation of DNA-PK in DU 145 cells

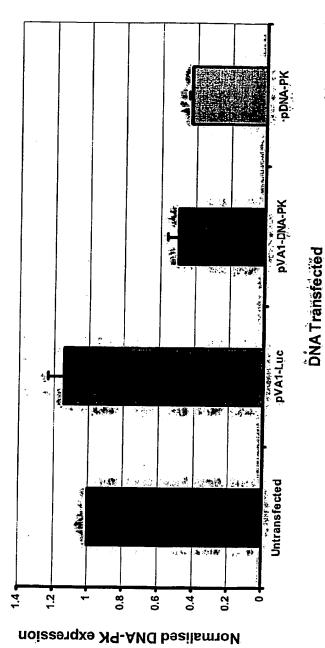
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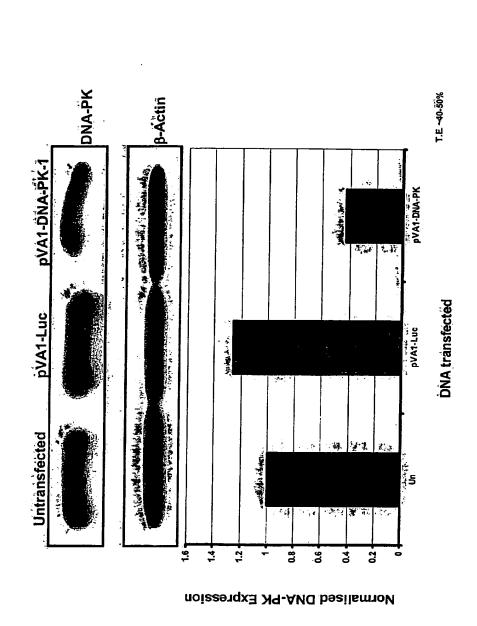
VA1 versus U6-mediated down-regulation of DNA-PK in DU 145 cells

DNA-PK expression 72hrs post-transfection



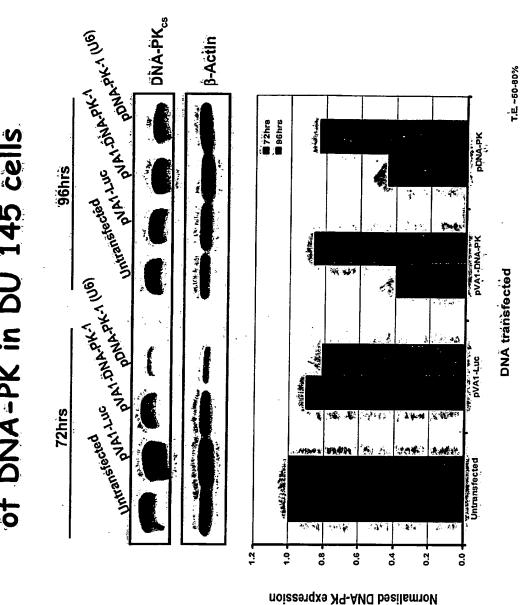
Average T.E ~50-60%

VA1-mediated döwn-regulation of DNA-PK in DU 145 cells (72hrs post-transfection)



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VA1 versus U6-mediated dówn-regulation of DNA-PK in DU 145 cells



### ATM-1 (5'- 3')

Oligo-A

TAGCTCTATCATGTTCTAGTTGACGGCAGAAGCTTGTGCCGTCGACTAGGACATGGTAGAGTTACAGTTTTTT

Oligo-B

GATCAAAAAACTGTAACTCTACCATGTCCTAGTCGACGGCACAAGCTTCTGCCGTCAACTAGAACATGATAGAGCTACG

Target sequence:-

TGCCGTCAACTAGAACATGATAGAGCTACAG (223-253, ATG = 190)

 $_{\circ}G/C$  content = 45%

ATM-2 (5'- 3')

Oligo-A

 ${\tt CCTGGAGGCTTGTGTGAGGCTGATACAGAGCTTGTGTATCAGCCTCAGCATAAGCCTCCGGGTAGTTTTTT}$ 

Oligo-B

GATCAAAAAACTACCCGGAGGCTTATGCTGAGGCTGATACACAAGCTTCTGTATCAGCCTCAACACAAGCCTCCAGGCG

Target sequence:-

TGTATCAGCCTCAACACACACACGCCTCCAGGCAG (432-462, ATG = 190)

G/C content = 55%

### ATM-3 (5'- 3')

### Oligo-A

TAGTATGTTGCTACAATCAGCTCCGTAAGAAGCTTGTTACGGAGCTGATTGTGGCGACGTATTACTCTTTTTT

### Oligo-B

GATCAAAAAAGAGTAATACGTCGCCACAATCAGCTCCGTAACAAGCTTCTTACGGAGCTGATTGTAGCAACATACTACG

### Target sequence:-

TTACGGAGCTGATTGTAGCAACATACTACTC (597-627, ATG = 190)

G/C content = 42%

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### ATR-1 (5'- 3')

### Oligo-A

TATTATATTCCTCTGGTGTGGCACTGCCGAAGCTTGGGCAGTGTCACACTAGAGGGATATAGTACAGTTTTTT

### Oligo-B

GATCAAAAAACTGTACTATATCCCTCTAGTGTGACACTGCCCAAGCTTCGGCAGTGCCACACCAGAGGAATATAATACG

### Target sequence:-

GGCAGTGCCACACCAGAGGAATATAATACAG (134-164, ATG = 80)

G/C content = 48%



### ATR-2 (5'- 3')

### Oligo-A

TTGCTGCAATCCGCAGAAGTCTCGTTATGAAGCTTGATAATGAGACTTCTGCGGATTGTAGTAATTCTTTTTT

### Oligo-B

GATCAAAAAAGAATTACTACAATCCGCAGAAGTCTCATTATCAAGCTTCATAACGAGACTTCTGCGGATTGCAGCAACG

### Target sequence:-

ATAACGAGACTTCTGCGGATTGCAGCAACC (388-418, ATG = 80)

G/C content = 48%

### ATR-3 (5'- 3')

Oligo-A

 $\tt CTCATGACCACTGGCCATTCCACAGCATGAAGCTTGATGCTGTGGAGTGGCCGGTGGTTATGAGTCGTTTTTT$ 

Oligo-B

GATCAAAAAACGACTCATAACCACCGGCCACTCCACAGCATCAAGCTTCATGCTGTGGAATGGCCAGTGGTCATGAGCG

Target sequence:-

ATGCTGTGGAATGGCCAGTGGTCATGAGCCG (579-609, ATG = 80)

G/C content = 58%



### **DNA-PK-1 (5'- 3')**

### Oligo-A

ATGTCTGTAATGCCAGCACCGCGGGGCTGAAGCTTGAGCCTCGTGGTGCTGGTATTACAGATATCTTTTTTT

### Oligo-B

GATCAAAAAAAAGATATCTGTAATACCAGCACCACGAGGCTCAAGCTTCAGCCCCGCGGTGCTGGCATTACAGACATCG

### Target sequence:-

AGCCCCGCGGTGCTGGCATTACAGACATCTT (196-226, ATG = 58)

G/C content = 58%

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## DNA-PK-2 (5'- 3')

Oligo-A

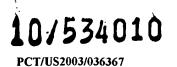
GATGAACTTCACCCAATAATCCTAGGAGGAAGCTTGCTTCTAGGATTATTGGGTGGAGTTCGTCTTATTTTTT

Oligo-B

Target sequence:-

CTCCTAGGATTATTGGGTGAAGTTCATCCTA (585-616, ATG = 58)

G/C content = 42%



### **DNA-PK-3 (5'-3')**

Oligo-A

TGAAGTTGCACAGAAGTGAGGACAACCCGAAGCTTGGGGTTGTTCTTACTTCTGTGCAGCTTCATTATTTTT

Oligo-B

GATCAAAAAATAATGAAGCTGCACAGAAGTAAGAACAACCCCAAGCTTCGGGTTGTCCTCACTTCTGTGCAACTTCACG

Target sequence:-

GGGTTGTCCTCACTTCTGTGCAACTTCACTA (733-763, ATG = 58)

G/C content = 48%

### ATM (5'- 3')

TAGCTCTATCATGTTCTAGTTGACGGCAX,TGCCGTCGACTAGGACATGGTAGAGTTACAGTTTTTT CCTGGAGGCTTGTGTTGAGGCTGATACAX,TGTATCAGCCTCAGCATAAGCCTCCGGGTAGTTTTT TAGTATGTTGCTACAATCAGCTCCGTAAX,TTACGGAGCTGATTGTGGCGACGTATTACTCTTTTTT

### ATR (5'- 3')

TATTATATTCCTCTGGTGTGGCACTGCCX,GGCAGTGTCACACTAGAGGGATATAGTACAGTTTTTT TTGCTGCAATCCGCAGAAGTCTCGTTATX,ATAATGAGACTTCTGCGGATTGTAGTAATTCTTTTT CTCATGACCACTGGCCATTCCACAGCATXATGCTGTGGAGTGGCCGGTGGTTATGAGTCGTTTTTT

### DNA-PK (5'- 3')

ATGTCTGTAATGCCAGCACCGCGGGGCTX,AGCCTCGTGGTGCTGGTATTACAGATATCTTTTTTT GATGAACTTCACCCAATAATCCTAGGAGX,CTTCTAGGATTATTGGGTGGAGTTCGTCTTATTTTTT TGAAGTTGCACAGAAGTGAGGACAACCCX,GGGTTGTTCTTACTTCTGTGCAGCTTCATTATTTTTT

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## Ad5 VA-1 (Top strand) - Accession No. X02996

10555 ctctgg

10561 ccggtcaggc gcgcgcaatc gttgacgctc tagaccgtgc aaaaggagag cctgtaagcg

10621 ggcactctt

10681 togagecceg tateeggeeg teegeegtga teeatgeggt taeegeegge gtg

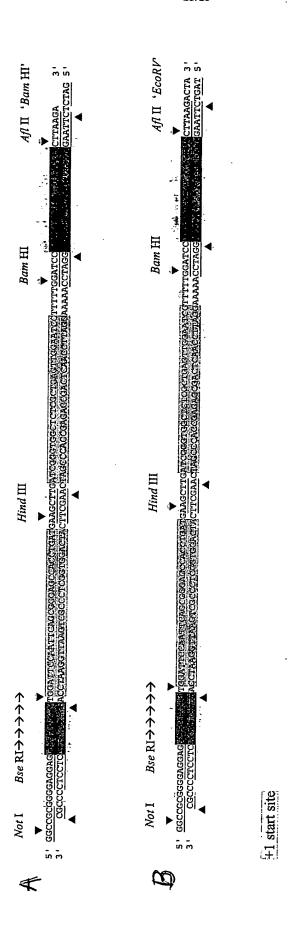
Variable upstream region

+1 start site

為建學的

B-Box





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